

Amendments to the Claims:

The following listing of claims will replace all prior versions, and listings, of claims in this application.

Listing of Claims

1. (Currently amended) A method for ~~treatment of~~ treating a subject having an apoptosis-related disease ~~in a subject~~ comprising administering to said subject a therapeutically effective amount of an inhibitor of ~~the~~ an isocitrate dehydrogenase (IDH) polypeptide, ~~wherein the IDH polypeptide~~ has an amino acid sequence as set forth in SEQ ID NO:2 or SEQ ID NO:4 or has a sequence which is modified therefrom while retaining the biological properties of IDH, in a dosage sufficient to inhibit expression of the IDH polypeptide, so as to thereby treat the subject.
2. (Currently amended) A method according to claim 1, wherein the inhibitor is administered in conjunction with a chemotherapeutic agent.
3. (Withdrawn- currently amended) A method according to claim 1, wherein the inhibitor is an antibody.
4. (Withdrawn- currently amended) A method according to claim 1, wherein the inhibitor is a chemical molecule selected from the group consisting of 2-(4-bromo-2,3-dioxobutylthio)-1, N6-ethenoadenosine 2',5'-bisphosphate, NADP oxoglutarate, o-(carboxymethyl) oxalohydroxamate, oxalylglycine, 3-bromo-2-ketoglutarate, beta-mercapto-alpha-ketoglutarate, beta-methylmercapto-alpha-ketoglutarate, beta-methylmercapto-

alpha-hydroxyglutarate, adriamycin and alpha-methylisocitrate.

5. (Withdrawn- currently amended) A method according to claim 1, wherein the inhibitor is an AS fragment comprising consecutive nucleotides having ~~the~~ a sequence as set forth in SEQ ID NO:5.
6. (Currently amended) A method according to claim 1, wherein the apoptosis-related disease is a cancer.
7. (Currently amended) A method for potentiating a chemotherapeutic treatment of a subject having an apoptosis-related disease ~~in a subject~~ comprising administering to said subject a therapeutically effective amount of an inhibitor of ~~the~~ a human isocitrate dehydrogenase (IDH) polypeptide in conjunction with a chemotherapeutic agent so as to thereby treat the subject; wherein the IDH polypeptide has an amino acid sequence as set forth in SEQ ID NO:2 or SEQ ID NO:4 or has a sequence which is modified therefrom while retaining the biological properties of IDH.
8. (Withdrawn- currently amended) A method according to claim 7, wherein the inhibitor is an antibody.
9. (Withdrawn- currently amended) A method according to claim 7, wherein the inhibitor is a chemical molecule selected from the group consisting of 2-(4-bromo-2,3-dioxobutylthio)-1, N6-ethenoadenosine 2',5'-bisphosphate, NADP oxoglutarate, o-(carboxymethyl) oxalohydroxamate, oxalylglycine, 3-bromo-2-ketoglutarate, beta-mercapto-alpha-ketoglutarate, beta-methylmercapto-alpha-ketoglutarate, beta-methylmercapto-alpha-hydroxyglutarate, adriamycin and alpha-

methylisocitrate.

10. (Withdrawn- currently amended) A method according to claim 7, wherein the inhibitor is an AS fragment comprising consecutive nucleotides having ~~the~~ a sequence as set forth in SEQ ID NO:5.
11. (Currently amended) A method according to claim 7, wherein the apoptosis-related disease is a cancer.
12. (Withdrawn- currently amended) An antisense oligonucleotide capable of inhibiting the expression of the IDH polypeptide, having ~~the~~ a sequence as set forth in SEQ ID NO:5.
13. (Withdrawn- currently amended) An expression vector comprising a nucleic acid ~~molecule~~ encoding the antisense oligonucleotide of claim 12.
14. (Withdrawn- currently amended) A process for determining the susceptibility of a subject to a chemotherapeutic treatment of an apoptosis-related disease comprising:
 - (a) providing ~~the~~ an average, normal level of the IDH polypeptide in ~~the~~ cells of healthy subjects;
 - (b) determining the level of the IDH polypeptide in said subject; and
 - (c) comparing the levels obtained in (a) and in (b) above, a low level of IDH polypeptide in said subject as compared to the level in healthy subjects indicating a susceptibility of said subject to a the chemotherapeutic treatment of said apoptosis-related disease.
15. (Withdrawn- currently amended) A process for determining the susceptibility of a subject to a chemotherapeutic

treatment of an apoptosis-related disease comprising:

- (a) providing ~~the~~ an average, normal level of mRNA encoding the IDH polypeptide in ~~the~~ cells of healthy subjects;
- (b) determining the level of mRNA encoding the IDH polypeptide in said subject; and
- (c) comparing the levels obtained in (a) and in (b) above, a low level of mRNA encoding IDH in said subject as compared to the level in healthy subjects indicating a susceptibility of said subject to a the chemotherapeutic treatment of said apoptosis-related disease.

16. (Withdrawn- currently amended) A process for determining the efficacy of a chemotherapeutic treatment administered to a subject comprising:

- (a) determining the level of the IDH polypeptide in the subject prior to a treatment;
- (b) determining the level of the IDH polypeptide in the subject after the treatment; and
- (c) comparing the levels obtained in (a) and in (b) above, a high level of IDH polypeptide prior to the treatment as compared to the level after the treatment indicating efficacy of the treatment.

17. (Withdrawn- currently amended) A process for determining the efficacy of a chemotherapeutic treatment administered to a subject comprising:

- (a) determining the level of ~~the~~ mRNA encoding an IDH mRNA polypeptide in the subject prior to a treatment;
- (b) determining the level of the IDH-encoding mRNA in the subject after the treatment; and
- (c) comparing the levels obtained in (a) and in (b) above, a high level of IDH-encoding mRNA prior to the treatment as compared to the level after the treatment indicating

efficacy of the treatment.

18. (Withdrawn) A process of diagnosing a cancer in a subject comprising:
- (a) providing ~~the~~ an average, normal level of the IDH polypeptide in ~~the~~ cells of healthy subjects;
 - (b) determining the level of the polypeptide in said subject; and
 - (c) comparing the levels obtained in (a) and in (b) above, wherein a high level of the IDH polypeptide in said subject as compared to the level in healthy subjects is indicative of a the cancer.
19. (Withdrawn- currently amended) A process of diagnosing a cancer in a subject comprising:
- (a) providing ~~the~~ an average, normal level of a polynucleotide encoding the IDH polypeptide in ~~the~~ cells of healthy subjects;
 - (b) determining the level of the polynucleotide in said subject; and
 - (c) comparing the levels obtained in (a) and in (b) above, wherein a high level of the polynucleotide in said subject as compared to the level in healthy subjects is indicative of a the cancer.
20. (Withdrawn) A process for obtaining a compound which modulates apoptosis in a cell comprising:
- (a) providing cells which express the human IDH polypeptide;
 - (b) contacting said cells with said compound; and
 - (c) determining the ability of said compound to modulate apoptosis in the cells.

21. (Withdrawn) A process according to claim 20 comprising:
- (a) providing test cells and control cells which express the human IDH polypeptide at a level at which approximately 50% of the cells undergo apoptosis in the presence of an apoptosis-stimulating agent;
 - (b) contacting said test cells with said compound;
 - (c) treating said cells in conjunction with step (b) with an amount of apoptosis-stimulating agent capable of causing apoptosis in the control cell; and
 - (d) determining the ability of said compound to modulate apoptosis in the test cell.
22. (Withdrawn) A process for obtaining a compound which promotes apoptosis in a cell comprising:
- (a) providing a test cell which expresses the human IDH polypeptide and a control cell which does not express the human IDH polypeptide;
 - (b) contacting said cells with said compound;
 - (c) treating said cells in conjunction with step (b) with an amount of apoptosis-stimulating agent capable of causing apoptosis in the control cell but not in the test cell in the absence of said compound; and
 - (d) determining the ability of said compound to promote apoptosis in the test cell.
23. (Withdrawn) A process for obtaining a compound which modulates apoptosis through the human IDH polypeptide comprising:
- (a) measuring the activity of the human IDH polypeptide, or a fragment thereof having viability activity,
 - (b) contacting said polypeptide or fragment with said compound; and
 - (c) determining whether the activity of said polypeptide or

fragment is modulated by said compound.

24. (Withdrawn) A process for obtaining a compound which modulates apoptosis through the human IDH polypeptide comprising:
- (a) measuring the binding of the human IDH polypeptide, or a fragment thereof having viability activity, to a species to which the human IDH polypeptide interacts specifically in vivo to produce an anti-apoptotic effect;
 - (b) contacting said polypeptide or fragment with said compound; and
 - (c) determining whether the activity of said polypeptide or fragment is affected by said compound.
25. (Currently amended) The method according to claim 1, wherein the inhibitor is an siRNA for ~~the~~ an IDH gene encoding an IDH polypeptide having an amino acid sequence as set forth in SEQ ID NO:2 or SEQ ID NO:4 or having a sequence which is modified therefrom while retaining the biological properties of IDH.
26. (Currently amended) The method according to claim 7, wherein the inhibitor is an siRNA for ~~the~~ an IDH gene encoding an IDH polypeptide having an amino acid sequence as set forth in SEQ ID NO:2 or SEQ ID NO:4 or having a sequence which is modified therefrom while retaining the biological properties of IDH.
27. (New) The method according to claim 25, wherein the inhibitor is an siRNA comprising nucleotides having a nucleotide sequence as set forth in SEQ ID NO:6.
28. (New) The method according to claim 26, wherein the

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inhibitor is an siRNA comprising nucleotides having a
nucleotide sequence as set forth in SEQ ID NO:6.